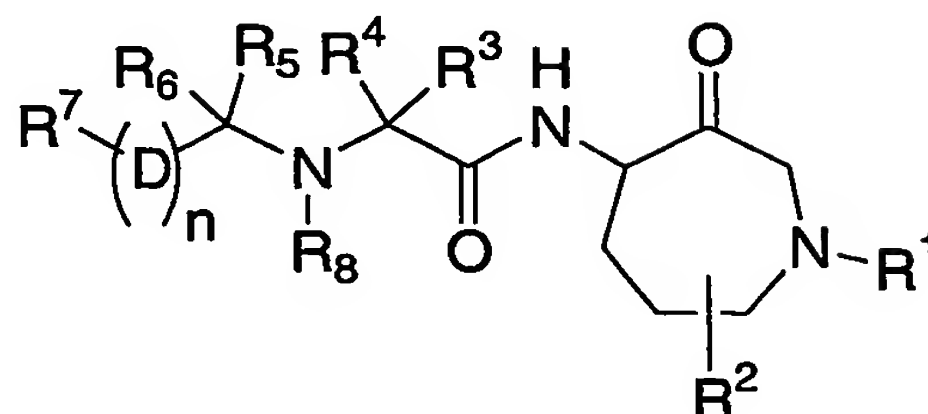


## WHAT IS CLAIMED IS:

1. A compound of the formula:



5 wherein R<sup>1</sup> is hydrogen, C<sub>1</sub>-6 alkyl, -SO<sub>2</sub>R<sup>9</sup>, -C(O)R<sup>9</sup> or arylC<sub>1</sub>-6alkyl;

R<sup>2</sup> is hydrogen, C<sub>1</sub>-6 alkyl or C<sub>3</sub>-6 cycloalkyl;

10 R<sup>3</sup> is hydrogen, C<sub>1</sub>-6 alkyl or C<sub>2</sub>-6 alkenyl wherein said alkyl and alkenyl groups are optionally substituted with C<sub>3</sub>-6 cycloalkyl or halo;

R<sup>4</sup> is hydrogen, C<sub>1</sub>-6 alkyl or C<sub>2</sub>-6 alkenyl wherein said alkyl and alkenyl groups are optionally substituted with C<sub>3</sub>-6 cycloalkyl or halo;

15 or R<sup>3</sup> and R<sup>4</sup> can be taken together with the carbon atom to which they are attached to form a C<sub>3</sub>-8 cycloalkyl ring, C<sub>5</sub>-8 cycloalkenyl ring, or five to seven membered heterocyclyl wherein said cycloalkyl, cycloalkenyl and heterocyclyl groups are optionally substituted with C<sub>1</sub>-6 alkyl, halo, hydroxyalkyl, hydroxy, alkoxy or keto;

R<sup>5</sup> is selected from hydrogen or C<sub>1</sub>-6 alkyl substituted with 1-6 halo;

20 R<sup>6</sup> is aryl, heteroaryl, C<sub>1</sub>-6 haloalkyl, arylalkyl or heteroarylalkyl, wherein said aryl, heteroaryl, arylalkyl and heteroarylalkyl groups are optionally substituted with halo, C<sub>1</sub>-6 alkyl, C<sub>1</sub>-6 haloalkyl, C<sub>3</sub>-6 cycloalkyl, -SR<sup>9</sup>, -SR<sup>12</sup>, -SOR<sup>9</sup>, -SOR<sup>12</sup>, -SO<sub>2</sub>R<sup>9</sup>, -SO<sub>2</sub>R<sup>12</sup>, -SO<sub>2</sub>CH(R<sup>12</sup>)(R<sup>11</sup>), -OR<sup>12</sup>, -N(R<sup>10</sup>)(R<sup>11</sup>) or cyano;

25 D is C<sub>1</sub>-3 alkyl, C<sub>2</sub>-3 alkenyl, C<sub>2</sub>-3 alkenyl, aryl, heteroaryl, C<sub>3</sub>-8 cycloalkyl or heterocyclyl wherein said aryl, heteroaryl, cycloalkyl and heterocyclyl groups, which may be monocyclic or bicyclic, are optionally substituted on either the carbon or the heteroatom with one to five substituents selected from C<sub>1</sub>-6 alkyl, halo or keto;

$R^7$  is hydrogen, C<sub>1-6</sub> alkyl, C<sub>2-6</sub> alkenyl, C<sub>2-6</sub> alkynyl, C<sub>1-6</sub> alkyloxy, halo, nitro, cyano, aryl, heteroaryl, C<sub>3-8</sub> cycloalkyl, heterocyclyl, -C(O)OR<sup>10</sup>, -C(O)OSi[CH(CH<sub>3</sub>)<sub>2</sub>]<sub>3</sub>, -OR<sup>10</sup>, -C(O)R<sup>10</sup>, -R<sup>10</sup>C(O)R<sup>9</sup>, -C(O)R<sup>9</sup>, -C(O)N(R<sup>12</sup>)(R<sup>12</sup>), -C(O)N(R<sup>10</sup>)(R<sup>11</sup>), -C(R<sup>10</sup>)(R<sup>11</sup>)OH, -SR<sup>12</sup>, -SR<sup>9</sup>, -R<sup>10</sup>SR<sup>9</sup>, -R<sup>9</sup>, -C(R<sup>9</sup>)<sub>3</sub>, -C(R<sup>10</sup>)(R<sup>11</sup>)N(R<sup>9</sup>)<sub>2</sub>, -NR<sup>10</sup>C(O)NR<sup>10</sup>S(O)<sub>2</sub>R<sup>9</sup>, -SO<sub>2</sub>R<sup>12</sup>, -SO(R<sup>12</sup>), -SO<sub>2</sub>R<sup>9</sup>, -SO<sub>2</sub>N(R<sup>c</sup>)(R<sup>d</sup>), -SO<sub>2</sub>CH(R<sup>10</sup>)(R<sup>11</sup>), -SO<sub>2</sub>N(R<sup>10</sup>)C(O)(R<sup>12</sup>), -SO<sub>2</sub>(R<sup>10</sup>)C(O)N(R<sup>12</sup>)<sub>2</sub>, -OSO<sub>2</sub>R<sup>10</sup>, -N(R<sup>10</sup>)(R<sup>11</sup>), -N(R<sup>10</sup>)C(O)N(R<sup>10</sup>)(R<sup>9</sup>), -N(R<sup>10</sup>)C(O)R<sup>10</sup>, -N(R<sup>10</sup>)C(O)OR<sup>10</sup>, -N(R<sup>10</sup>)SO<sub>2</sub>(R<sup>10</sup>), -C(R<sup>10</sup>)(R<sup>11</sup>)NR<sup>10</sup>C(R<sup>10</sup>)(R<sup>11</sup>)R<sup>9</sup>, -C(R<sup>10</sup>)(R<sup>11</sup>)N(R<sup>10</sup>)R<sup>9</sup>, -C(R<sup>10</sup>)(R<sup>11</sup>)N(R<sup>10</sup>)(R<sup>11</sup>), -C(R<sup>10</sup>)(R<sup>11</sup>)SC(R<sup>10</sup>)(R<sup>11</sup>)R<sup>9</sup>, R<sup>10</sup>S-, -C(R<sup>a</sup>)(R<sup>b</sup>)NR<sup>a</sup>C(R<sup>a</sup>)(R<sup>b</sup>)<sub>2</sub>, -C(R<sup>a</sup>)(R<sup>b</sup>)N(R<sup>a</sup>)(R<sup>b</sup>), -C(R<sup>a</sup>)(R<sup>b</sup>)C(R<sup>a</sup>)(R<sup>b</sup>)N(R<sup>a</sup>)(R<sup>b</sup>), -C(O)C(R<sup>a</sup>)(R<sup>b</sup>)N(R<sup>a</sup>)(R<sup>b</sup>), -C(R<sup>a</sup>)(R<sup>b</sup>)N(R<sup>a</sup>)C(O)R<sup>9</sup>, -C(O)C(R<sup>a</sup>)(R<sup>b</sup>)S(R<sup>a</sup>)(R<sup>b</sup>) or C(R<sup>a</sup>)(R<sup>b</sup>)C(O)N(R<sup>a</sup>)(R<sup>b</sup>); wherein said groups are optionally substituted on either the carbon or the heteroatom with one to five substituents independently selected from C<sub>1-6</sub> alkyl, halo, keto, cyano, haloalkyl, hydroxyalkyl, -OR<sup>9</sup>, -O(aryl), -NO<sub>2</sub>, -NH<sub>2</sub>, -NHS(O)<sub>2</sub>R<sup>8</sup>, -R<sup>9</sup>SO<sub>2</sub>R<sup>12</sup>, SO<sub>2</sub>R<sup>12</sup>, SO(R<sup>12</sup>), SO<sub>2</sub>N(R<sup>c</sup>)(R<sup>d</sup>), SO<sub>2</sub>N(R<sup>10</sup>)C(O)(R<sup>12</sup>), -C(R<sup>10</sup>)(R<sup>11</sup>)N(R<sup>10</sup>)(R<sup>11</sup>), -C(R<sup>10</sup>)(R<sup>11</sup>)OH, -COOH, -C(R<sup>a</sup>)(R<sup>b</sup>)C(O)N(R<sup>a</sup>)(R<sup>b</sup>), -N(R<sup>10</sup>)C(R<sup>10</sup>)(R<sup>11</sup>), -NH(CH<sub>2</sub>)<sub>2</sub>OH, -NHC(O)OR<sup>10</sup>, Si(CH<sub>3</sub>)<sub>3</sub>, heterocycl, aryl or heteroaryl;

$R^8$  is hydrogen or C<sub>1-6</sub> alkyl;  
 or  $R^4$  and  $R^8$  or can be taken together with any of the atoms to which they may be attached or are between them to form a 4-10 membered heterocycl ring system wherein said ring system, which may be monocyclic or bicyclic, is optionally substituted with C<sub>1-6</sub> alkyl, halo, hydroxyalkyl, hydroxy, keto, OR<sup>10</sup>, SR<sup>10</sup> or N(R<sup>10</sup>)<sub>2</sub>;

$R^9$  is selected from the group consisting of hydrogen, aryl, aryl(C<sub>1-4</sub>) alkyl, heteroaryl, heteroaryl(C<sub>1-4</sub>)alkyl, C<sub>3-8</sub>cycloalkyl, C<sub>3-8</sub>cycloalkyl(C<sub>1-4</sub>)alkyl, and heterocycl(C<sub>1-4</sub>)alkyl wherein said groups can be optionally substituted with halo or alkoxy;

$R^{10}$  is hydrogen or C<sub>1-6</sub> alkyl

$R^{11}$  is hydrogen or C<sub>1-6</sub> alkyl;

$R^{12}$  is hydrogen or C<sub>1-6</sub> alkyl which is optionally substituted with halo, alkoxy, cyano, -NR<sup>10</sup> or -SR<sup>10</sup>;

$R^a$  is hydrogen, C<sub>1-6</sub> alkyl, (C<sub>1-6</sub> alkyl)aryl, (C<sub>1-6</sub> alkyl)hydroxyl, -O(C<sub>1-6</sub> alkyl), hydroxyl, halo, aryl, heteroaryl, C<sub>3-8</sub> cycloalkyl, heterocycl, wherein said alkyl, aryl, heteroaryl, C<sub>3-8</sub>

cycloalkyl and heterocyclyl can be optionally substituted on either the carbon or the heteroatom with C<sub>1-6</sub> alkyl or halo;

R<sup>b</sup> is hydrogen, C<sub>1-6</sub> alkyl, (C<sub>1-6</sub> alkyl)aryl, (C<sub>1-6</sub> alkyl)hydroxyl, alkoxy, hydroxyl, halo, aryl, heteroaryl, C<sub>3-8</sub> cycloalkyl, heterocyclyl, wherein said alkyl, aryl, heteroaryl, C<sub>3-8</sub> cycloalkyl and heterocyclyl can be optionally substituted on either the carbon or the heteroatom with C<sub>1-6</sub> alkyl or halo;

or R<sup>a</sup> and R<sup>b</sup> can be taken together with the carbon atom to which they are attached or are between them to form a C<sub>3-8</sub> cycloalkyl ring or C<sub>3-8</sub> heterocyclyl ring wherein said 3-8 membered ring system may be optionally substituted with C<sub>1-6</sub> alkyl and halo;

R<sup>c</sup> is hydrogen or C<sub>1-6</sub> alkyl which is optionally substituted with halo or OR<sup>9</sup>;

R<sup>d</sup> is hydrogen or C<sub>1-6</sub> alkyl which is optionally substituted with halo or OR<sup>9</sup>;

or R<sup>c</sup> and R<sup>d</sup> can be taken together with the nitrogen atom to which they are attached or are between them to form a C<sub>3-8</sub> heterocyclyl ring which is optionally substituted with C<sub>1-6</sub> alkyl, halo hydroxyalkyl, hydroxy, alkoxy or keto;

n is independently selected from an integer from zero to three;

and the pharmaceutically acceptable salts, stereoisomers and N-oxide derivatives thereof.

2. The compound of Claim 2 wherein R<sup>3</sup> and R<sup>4</sup> are each independently selected from hydrogen or C<sub>1-4</sub> alkyl; or R<sup>3</sup> and R<sup>4</sup> can be taken together with the carbon atom to which they are attached to form a six membered cycloalkyl ring system, and the pharmaceutically acceptable salts, stereoisomers and N-oxide derivatives thereof.

3. The compound of Claim 1 wherein R<sup>5</sup> is C<sub>1-6</sub>alkyl substituted with 1-6 halo and R<sup>6</sup> is C<sub>1-6</sub> alkyl substituted with 1-6 halo; and the pharmaceutically acceptable salts, stereoisomers and N-oxide derivatives thereof.

4. The compound of Claim 1 wherein R<sup>5</sup> is hydrogen and R<sup>6</sup> is C<sub>1-6</sub> alkyl substituted with 1-6 halo; and the pharmaceutically acceptable salts, stereoisomers and N-oxide derivatives thereof.

5. The compound of Claim 1 wherein R<sup>5</sup> is hydrogen and R<sup>6</sup> is aryl or heteroaryl wherein said aryl or heteroaryl are optionally substituted with halo or -SO<sub>2</sub>R<sup>12</sup>; and the pharmaceutically acceptable salts, stereoisomers and N-oxide derivatives thereof.

5 6. The compound of Claim 1 wherein R<sup>4</sup> and R<sup>8</sup> or can be taken together with any of the atoms to which they may be attached or are between them to form a 4-10 membered heterocyclyl ring system wherein said ring system, which may be monocyclic or bicyclic, is optionally substituted with C<sub>1-6</sub> alkyl, halo, hydroxyalkyl, hydroxy, keto, -OR<sup>10</sup>, -SR<sup>10</sup> or -N(R<sup>10</sup>)<sub>2</sub>; and the pharmaceutically acceptable salts, stereoisomers and N-oxide  
10 derivatives thereof.

7. The compound of Claim 6 wherein R<sup>4</sup> and R<sup>8</sup> can be taken together with any of the atoms to which they may be attached or are between them to form a 5 or 6 membered heterocyclyl ring system wherein said ring system, is optionally substituted with C<sub>1-6</sub> alkyl, halo,  
15 hydroxyalkyl, hydroxy, keto, -OR<sup>10</sup>, -SR<sup>10</sup> or -N(R<sup>10</sup>)<sub>2</sub>; and the pharmaceutically acceptable salts, stereoisomers and N-oxide derivatives thereof.

8. The compound of Claim 1 selected from:  
*N*<sup>1</sup>-[3-Oxo-1-(pyridin-2-ylsulfonyl)azepan-4-yl]-*N*<sup>2</sup>-{(1*S*)-2,2,2-trifluoro-1-[4'-(methylsulfonyl)-  
20 1,1'-biphenyl-4-yl]ethyl}-L-leucinamide, and the pharmaceutically acceptable salts, stereoisomers and N-oxide derivatives thereof.

9. A pharmaceutical composition comprising a compound according to Claim 1 and a pharmaceutically acceptable carrier.

25 10. A pharmaceutical composition made by combining a compound according to any one of Claims 1 to 8 and a pharmaceutically acceptable carrier.

30 11. A process for making a pharmaceutical composition comprising combining a compound according to Claim 1 and a pharmaceutically acceptable carrier.

12. A method of inhibiting cathepsin activity in a mammal in need thereof, comprising administering to the mammal a therapeutically effective amount of a compound according to Claim 1.

13. The method according to Claim 13 wherein the cathepsin activity is Cathepsin K activity.

14. A method of treating or preventing a disease selected from: osteoporosis, glucocorticoid induced osteoporosis, Paget's disease, abnormally increased bone turnover, periodontal disease, tooth loss, bone fractures, rheumatoid arthritis, osteoarthritis, periprosthetic osteolysis, osteogenesis imperfecta, metastatic bone disease, hypercalcemia of malignancy or multiple myeloma in a mammal in need thereof by administering to the mammal a therapeutically effective amount of a compound according to Claim 1.

15. A method of treating or preventing bone loss in a mammal in need thereof by administering to the mammal a therapeutically effective amount of a compound according to Claim 1.

16. A method of treating or preventing osteoporosis in a mammal in need thereof by administering to the mammal a therapeutically effective amount of a compound according to Claim 1.

17. A method of treating cathepsin dependent conditions in a mammal in need thereof by administering to the mammal a therapeutically effective amount of a compound according to Claim 1.

18. A pharmaceutical composition comprising a compound of Claim 1 and another agent selected from: an organic bisphosphonate, an estrogen receptor modulator, an androgen receptor modulator, an inhibitor of osteoclast proton ATPase, an inhibitor of HMG-CoA reductase, an integrin receptor antagonist, or an osteoblast anabolic agent, and the pharmaceutically acceptable salts and mixtures thereof.

19. A method of treating osteoporosis comprising a compound of Claim 1 and another agent selected from: an organic bisphosphonate, an estrogen receptor modulator, an androgen receptor modulator, an inhibitor of osteoclast proton ATPase, an inhibitor of HMG-CoA reductase, an integrin receptor antagonist, or an osteoblast anabolic agent, and the pharmaceutically acceptable salts and mixtures thereof.

20. A method of treating bone loss comprising a compound of Claim 1 and another agent selected from: an organic bisphosphonate, an estrogen receptor modulator, an androgen receptor modulator, an inhibitor of osteoclast proton ATPase, an inhibitor of HMG-CoA reductase, an integrin receptor antagonist, or an osteoblast anabolic agent, and the pharmaceutically acceptable salts and mixtures thereof.

21. A pharmaceutical composition comprising a compound of Claim 1 and another agent selected from: an organic bisphosphonate, an estrogen receptor modulator, an androgen receptor modulator, an inhibitor of osteoclast proton ATPase, an inhibitor of HMG-CoA reductase, an integrin receptor antagonist, or an osteoblast anabolic agent, and the pharmaceutically acceptable salts and mixtures thereof.

22. A compound of any one of Claims 1 to 8, or a pharmaceutically acceptable salt, stereoisomer or N-oxide derivative thereof, for use in inhibiting cathepsin activity, such as cathepsin K activity.

23. Use of a compound of any one of Claims 1 to 8, or a pharmaceutically acceptable salt, stereoisomer or N-oxide derivative thereof, in the manufacture of a medicament for treating or preventing a disease set forth in Claim 14.